

CLAIMS

We claim:

- 1. A method of isolating one strand of a double-stranded target nucleic acid, comprising:
- (i) contacting a double-stranded target nucleic acid comprising a first strand and a second strand with a competitor oligo capable of hybridizing to the first strand under conditions in which the first strand dissociates from the second strand and hybridizes with the competitor oligo to form a first-strand:competitor oligo heterduplex; and
 - (ii) isolating the heteroduplex or the dissociated second strand.
- 2. A method of isolating one strand of a double-stranded target nucleic acid, comprising:
- (i) dissociating the double-stranded target nucleic acid into a first strand and a second strand;
- (ii) contacting the dissociated target nucleic acid with a competitor oligo capable of hybridizing to the first strand under conditions which favor first-strand:competitor oligo heteroduplex formation and disfavor reannealing of the first and second strands; and
 - (iii) isolating the heteroduplex or the dissociated second strand.
 - 3. The method of Claim 1 or 2 in which in step (ii) the heteroduplex is isolated.
- 4. The method of Claim 3 further comprising the step of dissociating the heteroduplex and isolating the first strand.
 - 5. The method of Claim 1 or 2 in which in step (ii) the second strand is isolated.
- 6. The method of Claim 1 or 2 in which the double-stranded target nucleic acid is a double-stranded DNA.



- 7. The method of Claim 1 or 2 in which the double-stranded target nucleic acid is a double-stranded DNA/RNA hybrid duplex.
- 8. The method of Claim 1 or 2 in which the competitor oligo is composed of between 7 and 40 nucleobases.
- 9. The method of Claim 1 or 2 in which the double-stranded target nucleic acid has the formula:

TAIL 1---SEQUENCE---TAIL 2

TAIL 1'--SEQUENCE'--TAIL 2'

wherein:

TAIL 1 represents a first tail nucleobase sequence;

SEQUENCE represents a target nucleobase sequence;

TAIL 2 represents a second tail nucleobase sequence;

TAIL 1' represents a nucleobase sequence that is complementary to

TAIL 1;

SEQUENCE' represents a nucleobase sequence that is complementary

to SEQUENCE; and

TAIL 2' represents a nucleobase sequence that is complementary to

TAIL 2.

10. The method of Claim 9 in which a portion of the competitor oligo is capable of hybridizing to TAIL 1 and another portion of the competitor oligo is capable of hybridizing to TAIL 2.



- 11. The method of Claim 9 in which a portion of the competitor oligo is capable of hybridizing to TAIL 1' and another portion of the competitor oligo is capable of hybridizing to TAIL 2'.
- 12. The method of Claim 9 in which TAIL 1 and TAIL 2 comprise non-standard synthetic nucleobases.
- 13. The method of Claim 9 in which TAIL 1 and TAIL 2 are not complementary to one another.
- 14. The method of Claim 1 or 2, in which the competitor oligo includes a capture moiety.
- 15. The method of Claim 14, in which the capture moiety is one member of a pair of molecules that specifically bind to each other.
 - 16. The method of Claim 15, in which the capture moiety is biotin.
 - 17. The method of Claim 14, in which the moiety is a solid support.
 - 18. The method of Claim 17, in which the solid support is magnetic.
 - 19. The method of Claim 14 in which the capture moiety is a capture sequence.
 - 20. The method of Claim 16 in which the capture moiety is a charged group.
- 21. The method of Claim 1 or 2 in which the competitor oligo is capable of hybridizing to only the first or the second strand of the double-stranded target nucleic acid.



- 22. The method of Claim 1 or 2 in which the contacting step is carried out at a cationic strength in the range of 0 to 10 mM, a pH in the range of 6 to 8, and a temperature in the range of 20 to 40°C.
- 23. The method of Claim 1 or 2 in which the competitor oligo is a PNA and optionally includes from 1 to 4 positively charged nucleobase interlinkages.
- 24. The method of Claim 1 or 2 in which the competitor oligo comprises non-standard synthetic nucleobases.
- 25. A method of isolating one strand of a double-stranded target nucleic acid, comprising the steps of:
- (i) dissociating the double-stranded target nucleic acid into a first strand and a second strand;
- (ii) contacting the dissociated target nucleic acid with a competitor oligo capable of hybridizing to only the first strand under conditions which kinetically favor competitor oligo first-strand hybrid formation and kinetically disfavor reannealing of the first and second strands, said competitor oligo being conjugated with a moiety that facilitates capture of competitor oligo:first-strand hybrids; and
 - (iii) capturing the competitor oligo:first strand hybrid.
 - 26. The method of Claim 25 wherein the competitor oligo is a PNA.